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NEWS 40 May 19 Simultaneous left and right truncation added to WSCA

right truncation

NEWS 41 May 19 FAPFA enhanced with new search field, simultaneous left and

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ANSWER 1 OF 5 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

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ACCESSION NUMBER: 2002:219032 BIOSIS DOCUMENT NUMBER: PREV200200219032

TITLE: Mammalian phospholipase A2 nucleotide sequences, low molecular weight amino acid sequences encoded thereby,

antisense sequences and nucleotide sequences having

internal ribosome kinding sites.

AUTHOR(S): Tischfield, Jay A. (1); Seilhamer, Jeffrey J.

CORPORATE SOURCE: (1) 9982 Mill Fun, Carmel, IN, 46032 USA

ASSIGNEE: Tischfield; Jay A., Piscataway, NJ, USA; Incyte

Pharmaceuticals, Inc.

PATENT INFOFMATION: US 6352849 March 05, 2002

Official Gazette of the United States Patent and Trademark SOURCE:

Office Patents, (Mar. 5, 2002) Vol. 1256, No. 1, pp. No Pagination. http://www.uspto.gov/web/menu/patdata.html.

e-file.

ISSN: 0098-1133.

DOCUMENT TYPE: Pater.t LANGUAGE: English

Novel mammalian phospholipase (PLA2) nucleotide sequences and low molecular weight (about 14 KD) amino acid sequences encoded thereby are disclosed. More particularly, a cloned human HPLA2 cDNA expressing

HPLA2 -10 and its cloned rat RPLA2 cDNA counterpart, expressing RPLA2 -10, which are characterized as PLA2 Type IV, are disclosed. A second type of PLA2 cDNA, characterized as PLA2 Type III and

exemplified by a rat PLA2 cDNA encoding RPLA2 -8 and a partial human PLA2 nucleotide sequence encoding HPLA2 -8, is disclosed. Expression of the cDNAs encode the two new types of PLA2 enzymes which have phospholipase activity. The nevel FLA2 s do not include disulfide bridges between cysteine amino acids 12 and 77 or elapid loops. However, the nevel PLA2 s may include amino acid COOH-terminal extensions which can vary in length. Seventeen of the eighteen absolutely conserved amino acids in all active 14 KD PLA2 s are believed to be conserved in RPLA2 -8 and HPLA2 -8, whereas all eighteen are believed to be conserved in RPLA2 -10 and

HPLA2 -10. Because the encoded sequences of EPLA2 -8 and HPLA2 -8 include only 16 cysteine amino acids, they have been designated

as Type III. RPLA2 -10 and HPLA2 -10 are designated as Type IV since their encoded sequences include only 12 cysteine amino acids.

ANSWER 2 OF 5 BIOSIS COFFFIGHT 2003 BIOLOGICAL ABSTRACTS INC.

ACCESSION NUMBER: 2000:278419 BIOSIS DOCUMENT NUMBER: PREV200000278419

TITLE: Mammalian phospholipase A2 nucleotide sequences low

molecular weight amino acid sequences encoded thereby antisense sequences and nucleotide sequences having

internal ribosome binding sites.

Tischfield, Jay A. (1); Seilhamer, Jeffrey J. AUTHOR(S):

CORPORATE SOURCE: (1) Lcs Altos Hills, CA USA

ASSIGNEE: Tischfield; J., J., USA; Incyte Pharmaceuticals,

Inc., USA

PATENT INFORMATION: US 5972677 October 26, 1999

SOURCE:

Official Gazette of the United States Patent and Trademark Office Patents, (Oct. 26, 1999) Vol. 1227, No. 4, pp. No.

> pagination. e-file. ISSN: 0098-1133.

DOCUMENT TYPE: Patent LANGUAGE: English

Novel mammalian phospholipase (PLA.:) nucleotide sequences and low molecular weight (about 14KD) amino acid sequences encoded thereby are disclosed. More particularly, a cloned human HPLA2 cPNA expressing HPLA2 -10 and its cloned rat F.PLA2 cDNA counterpart,

expressing RPLA2 -10, which are characterized as PLA2 Type IV, are disclosed. A second type of PLA2 cDNA, characterized as PLA2 Type III and

exemplified by a rat PLA2 cDNA encoding RPLA2 -8 and a partial human PLA2 nucleotide sequence encoding HPLA2 -8, is disclosed. Expression of the cDNAs encode the two new types of FLA2 enzymes which have phospholipase activity. The novel PLA2 s do not include disulfide bridges between cysteine amino acids 11 and 77 or elapid loops. However, the novel PLA2 s may include amino acid COOH-terminal extensions which can vary in length. Seventeen of the eighteen absolutely conserved amino acids in all active 14KD PLA2 s are believed to be conserved in RPLA2 -8 and HPLA2 -8, whereas all eighteen are believed to be conserved in RPLA2 -10 and HPLA2 -10. Because the encoded sequences of RPLA2 -8 and HPLA2 -8include only 16 cysteine amino acids, they have been designated as Type III. PPLA2 -10 and HPLA2 -10 are designated as Type IV since their encoded sequences include only 12 cysteine amino acids.

ANSWER 3 OF 5 CA COPYRIGHT 2003 ACS

ACCESSION NUMBER:

131:298644 CA

TITLE:

Group V phospholipase A2-dependent induction of

cyclooxygenase-2 in macrophages

AUTHOR(S):

Balsınde, Jesus; Shinohara, Hiroyuki; Lefkowitz, Lee J.; Johnson, Christina A.; Balbca, Maria A.; Dennis,

Edward A.

CORPORATE SOURCE:

Department of Chemistry and Brochemistry, University of California at San Diego, La Jolla, CA, 92093-0601,

SOURCE:

Journal of Biological Chemistry (1999), 274(37),

25967-25970

CODEN: JBCHA3; ISSN: 0021-9258

PUBLISHER:

American Society for Biochemistry and Molecular

Bielogy Journal

DOCUMENT TYPE:

English

LANGUAGE: When exposed for prolonged periods of time (up to $20\ h$) to bacterial lipopolysaccharide (LPS) murine P388D1 macrophages exhibit a delayed prostaglandin biosynthetic response that is entirely mediated by cyclooxygenase-2 (COM-1). Both the constitutive Group IV cytosolic phospholipase A2 (cPLA2) and the inducible Group V secretory phospholipase A2 (sPLA2) are involved in the cyclopxygenase-2-dependent generation of prostaglandins in response to LPS. Using the selective sPLA2 inhibitor 3-(3-acetamide-1-benzyl-2-ethylindolyl-5-oxy)propane sulfonic acid (LY311727) and an antisense oligonucleotide specific for Group V sPLA2, the authors found that induction of COW-2 expression is strikingly dependent on Group V sFLA2, which was further confirmed by expts. in which exogenous Group V sFLA2 was added to the cells. Exogenous Group V sPLA2 was able to induce arachidonate mobilization on its own and to induce expression of the CCM-2. None of these effects was obsd. if inactive Group V sPLA2 was utilized, implying that enzyme activity is crucial for these effects to take place. Therefore, not only delayed prostaglandin prodn. but also COX-2 gene induction are dependent on a catalytically active Group V sPLA2. COX-2 expression was also blunted by the Group IV cPLA2 inhibitor Me arachidonyl flucrophosphonate, which the authors have previously found to block Group V sFLA2 induction as well. Collectively, the results support a model whereby Group IV cFLA2 activation regulates the expression of Group V sPLA2, which in turn is responsible for delayed prostaglandin prodn. by regulating COX-2 expression.

REFERENCE COUNT: 3.0

THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 4 OF 5 CA COPYRIGHT 2003 ACS ACCESSION NUMBER:

127:79056 CA

TITLE:

Analysis of the secretory phospholipase A2 that mediates prostaglandin production in mast cells

AUTHOR(S):

Pedd;, Srinivasa T.; Winstead, Michelle V.; Tischfield, Jay A.; Herschman, Harvey R.

CORPORATE SCURCE: Departments Biological Chemistry Molecular Medical

Fharmacology and the Molecular Biology Institute, UCLA Center Health Sciences, Los Angeles, CA, 90095-1570,

USA

SOURCE: Journal of Biological Chemistry (1997), 272(21),

13591-13596

CODEN: JECHA3; ISSN: 0021-9258

FUBLISHER: American Society for Biochemistry and Molecular

Biology

DGCUMENT TYPE: Journal LANGUAGE: English

Prostaglandin D2 (PGD2) synthesis in activated mast cells occurs in two phases, an early phase that is dependent on prostaglandin synthase 1 and a delayed phase that is dependent on activation-induced prostaglandin synthase 2 gene expression. Early phase PGD2 synthesis in activated mast cells also requires the activity of a secretory phospholipase A2 (PLA2). It has been thought that the secretory PLA2 expressed in mast cells is group IIa PLA2, encoded by the Fla2 g2a gene. However, activated bone marrow-derived mast cells prepd. from Pla2 g2a+/+ mice and mast cells prepd. from mice with a mutation in the Pla2 g2a gene both demonstrate early phase PGD2 synthesis. Moreover, mast cells from both murine strains secrete FLA2 activity following activation. Northern and reverse transcriptase/polymerase chain reaction analyses demonstrate that mast cells from Pla2 g2a+/+ and Pla2 g2a-/- mice do not express group IIa PLA2 message. Instead, Northern and reverse transcriptase/polymerase chain reaction analyses demonstrate that both Pla2 g2a+/+ and Pla2 g2a-/- mast cells express mFNA for group V FLA2, encoded by the Pla2qV gene. An antisense oligonuclectide directed against group V PLA2 is also able to inhibit both the early phase pf FGD2 prodn. and the secretion of PLA2 activity by activated mast cells. Our data suggest that (i) group IIa PLA2 does not play a significant role in murine mast cell prostaglandin synthesis, (ii) group V PLA2 mediates early mast cell PGD2 prodn. and transcellular PGE2 prodn. in murine mast cells, and (iii) much of the data, based on studies with chem. inhibitors and antibodies, suggesting that group IIa PLA2 is responsible for arachidonic acid mobilization needs to be reevaluated.

L4 ANSWER 5 OF 5 CA COPYRIGHT 2003 ACS ACCESSION NUMBER: 122:308082 CA

TITLE:

E: Mammalian low molecular weight phospholipase A2

nucleotide and amino acid sequences

INVENTOR(S): Tischfield, Jay A.; Seilhamer, Jeffrey J. PATENT ASSIGNEE(S): Indiana University Foundation, USA; Incyte

Pharmaceuticals, Inc. PCT Int. Appl., 159 pp.

CODEN: PIMMD2

DOCUMENT TYPE: Patent

LANGUAGE: Fatent English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

SOURCE:

PATENT NO. K				KI	KIND DATE			APPLICATION NO.					DATE				
WO	WO 9502328			A1 19950106				WO 1994-US7926 19940715									
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CA 2167296		AA 19950126															
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	AU 9473622							AU 1994-73622 19940715									
US 5972677			A 1999102		1026		US 1997-888497			7	19970707						

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PRIORITY APPLN. INFO.:
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                                        US 1993-97354
                                                        A 19930726
                                        WO 1994-US7926 W 19940715
                                        US 1996-651405 B1 19960522
                                        US 1997-888497 A3 19970707
AB
     Novel mammalian phospholipase (PLA2) nucleotide sequences and low mol. wt.
     (about 14 KD) amino acid sequences encoded thereby are disclosed. More
     particularly, a cloned human HPLA2 cDNA expressing HPLA2-
     10 and its cloned rat RPLA2 cDNA counterpart, expressing RPLA2-10,
     which are characterized as PLA2 Type IV, are disclosed. A second type of
     PLA2 cDNA, characterized as PLA2 Type III and exemplified by a rat PLA2
     cDNA encoding RPLA2-8 and a partial human FLA2 nucleotide sequence
     encoding HPLA2-3, is disclosed. Expression f the cDNAs encode the two new
     types of PLA2 enzymes which have phospholipase activity. The novel PLA2s
     do not include disulfide bridges between cysteine amino acids 11 and 77 or
     elapid loops. However, the novel PLA2s may include amino acid
     COOH-terminal extensions which can vary in length. Seventeen of the
     eighteen absolutely conserved amino acids in all active 14 KD PLA2s are
     believed to be conserved in PPLA2-3 and HPLA2-8, whereas all eighteen are
     believed to be conserved in EPLA2-10 and HPLA2-10.
     Because the encoded sequences of EPLA2-8 and HPLA2-8 include only 16
     cysteine amino acids, they have been designated as Type III. RPLA2-10 and
     HPLA2-10 are designated as Type IV since their encoded
     sequences include only 12 cysteine amino acids.
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